

## How to adjust drug dosing after bariatric surgery

**B**ariatric surgery has increased 10-fold in the United States over the past decade.<sup>1</sup> Because one-third of bariatric candidates take psychiatric medications, psychiatric input on postoperative management is in demand.<sup>2</sup>

Despite this surgery's rising popularity, no data exist to guide psychotropic dosing in patients with altered GI environments and who experience massive postoperative weight loss. Evidence and clinical experience support these recommendations:

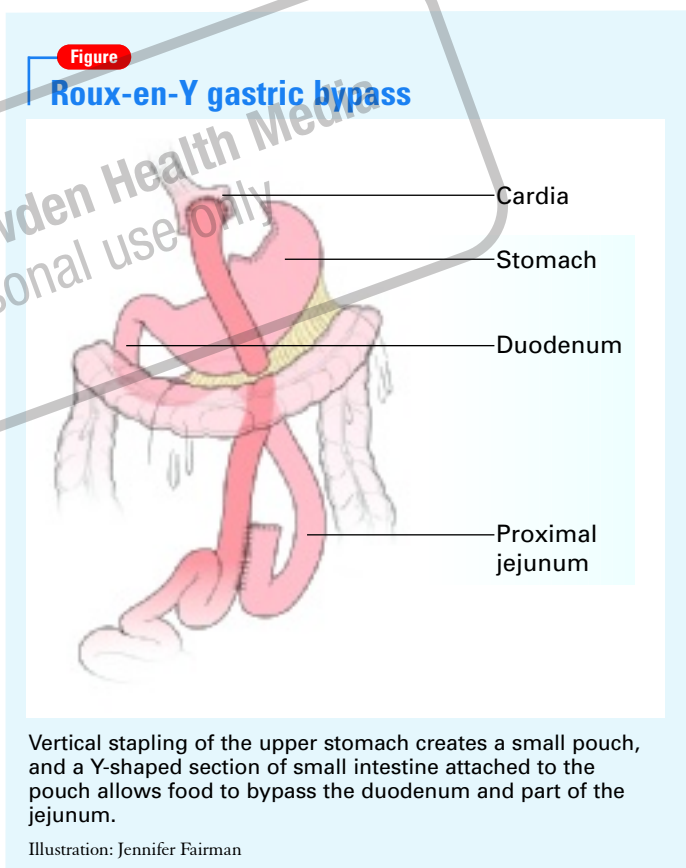
### Managing changes in psychotropic absorption.

The most common procedure, the Roux-en-Y gastric bypass (RYGB), bypasses the duodenum, proximal jejunum, and all of the stomach except the cardia (*Figure*). Because ingested food is normally absorbed in the duodenum and jejunum, a bypass results in relative malabsorption of food. Orally ingested food and medications pass into the cardia, where they are not exposed to acid.

One novel in vitro study<sup>2</sup> examined the dissolution of 22 psychotropics in a simulated GI environment of control and post-RYGB states. Twelve medications dissolved differently in a postoperative environment than in the control state; 10 dissolved much less than expected.

The effect of bariatric surgery on in vivo absorption of commonly used psychotropics has not been studied. For safe yet effective dosing immediately after surgery, try:

- **Using immediate-release psychotropics.**<sup>3</sup> Time-released medications are designed to dissolve gradual-



ly within the full intestinal tract. Extended-release formulations will probably have markedly different pharmacokinetics after RYGB. Coated pills require stomach acid to release active ingredients, so crush oral medications in the immediate postoperative period.

- **Monitoring serum drug levels of medications with a narrow therapeutic index.** Also measure preoperative blood levels while the patient feels well. This

Table

Common drugs' volume of distribution ( $V_d$ )

Drug	Extent of distribution	Volume of distribution
<b>Fluoxetine</b>	Large	10 to 103 L/kg
<b>Citalopram</b>	Moderate	14 to 17 L/kg
<b>Venlafaxine</b>	Moderate	6 to 7 L/kg
<b>Oxcarbazepine</b>	Small	0.7 L/kg
<b>Lithium</b>	Small	0.4 to 0.6 L/kg
<b>Valproic acid</b>	Small	0.1 to 0.4 L/kg

baseline will provide a target to aim for if the patient has a postoperative relapse.

**Postoperative weight loss.** Many physiologic factors influence drug absorption, such as gastric emptying time and the integrity and surface area of the epithelium. The gut has an impressive capacity to compensate for loss of function, so absorption after surgery may eventually normalize. By that time, however, marked weight loss can complicate the clinical picture.

Compared with nonobese persons, obese persons have an increased proportion of adipose tissue, as well as increased total body water, lean body mass, visceral organ mass, and higher glomerular filtration rate.

Postoperative bariatric patients often lose more than 100 pounds of adipose tissue. This type of weight loss mostly affects lipid-soluble

drugs with a large volume of distribution ( $V_d$ ) that readily cross cell membranes, such as fluoxetine and oxcarbazepine. Drugs with a large  $V_d$ , such as fluoxetine (Table), reach all major compartments of distribution, which in a normal weight individual include: plasma (5%), interstitial fluid (16%), intracellular fluid (35%), transcellular fluid (2%), and fat (20%). Because the amount of fat in an overweight patient is initially very high but rapidly decreases after bariatric surgery, drugs with a large  $V_d$  can shift into other compartments.

For drugs with a small  $V_d$ , such as lithium, a lower maintenance dosage may be required because of decreased glomerular filtration following marked weight loss.<sup>4</sup>

References

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